

[illegible]

```

As.6.1      EIQLVQSGPELKQPGETVRISCKASGYTFTNYGMNVWKQAPGKGLKWMG
              *  *  *      *  *  *      *  *  *
E(-)-12     EVQLVESGGGLVQPGGSLRLSCAASGYTFTNYGMNVWRQAPGKGLEWVG
              *  *  *  *
HumIII      EVQLVESGGGLVQPGGSLRLSCAASGFTFSSYAMSWVRQAPGKGLEWVS
              1      10      20      30      40

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A4.6.1      WINTYTGEPTYAADEKRRFTFSLETSASTAYLQISNLKNDDETATYFCAK
              * * * * *
F(ab)-12    WINTYTGEPTYAADEKRRFTFSLDTSKSTAYLQMNSLRAEDTAVYYCAK
              * * * * *
humIII      VISGDGGSTYYADSVKGRFTISRDN SKNTLYLQMNSLRAEDTAVYYCAR
              50 a          60          70          80 abc          90

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Fig. 1A

A.4.6.1 YPHYYGSSSHWYFDVWGAGTTVTVSS (SEQ ID NO:9)
 * *
F(ab)-12 YPHYYGSSSHWYFDVWGQGTLVTVSS (SEQ ID NO:7)
 * *
humIII G-----FDYWGQGTLVTVSS (SEQ ID NO:11)

110

Variable Light

A4.6.1 DIQMTQTSSLSASLGDRVIISCSASODISNYLNWYQQKPDGTVKVLIY
 ** * * * ****

F(ab)-12 DIQMTQSPSSLSASVGDRVITICSASODISNYLNWYQQKPGKAPKVLII
 * * * *

humKI DIQMTQSPSSLSASVGDRVITICRASQSISNYLAWYQQKFSGKAPKLLIY
 1 10 20 30 40

Fig. 1B

A4.6.1 FTSSLHSGVPSRFSGSGSGTDYSLTISNLEPEDIATYYCQOYSTVPWTF
 ** * *

F(ab)-12 FTESL^HS^LGVPSPSRFSGSGSGTDFTLTISSLQPEDFATYYCQOYSTVF^TWF^T
 ** * ***

humKI AASSLESGLVPSRFSGSGSGTDFTLTISSLQPEDFATYYCQQYNLPWTF
30 60 70 80 90

A4.6.1 GGGTKI^{*}EIKR (SEQ ID NO: 10)
F(ab)-12 GQGTKV^{*}EIKR (SEQ ID NO: 8)
humKI GQGTKV^{*}EIKR (SEQ ID NO: 12)
100

08/908,469

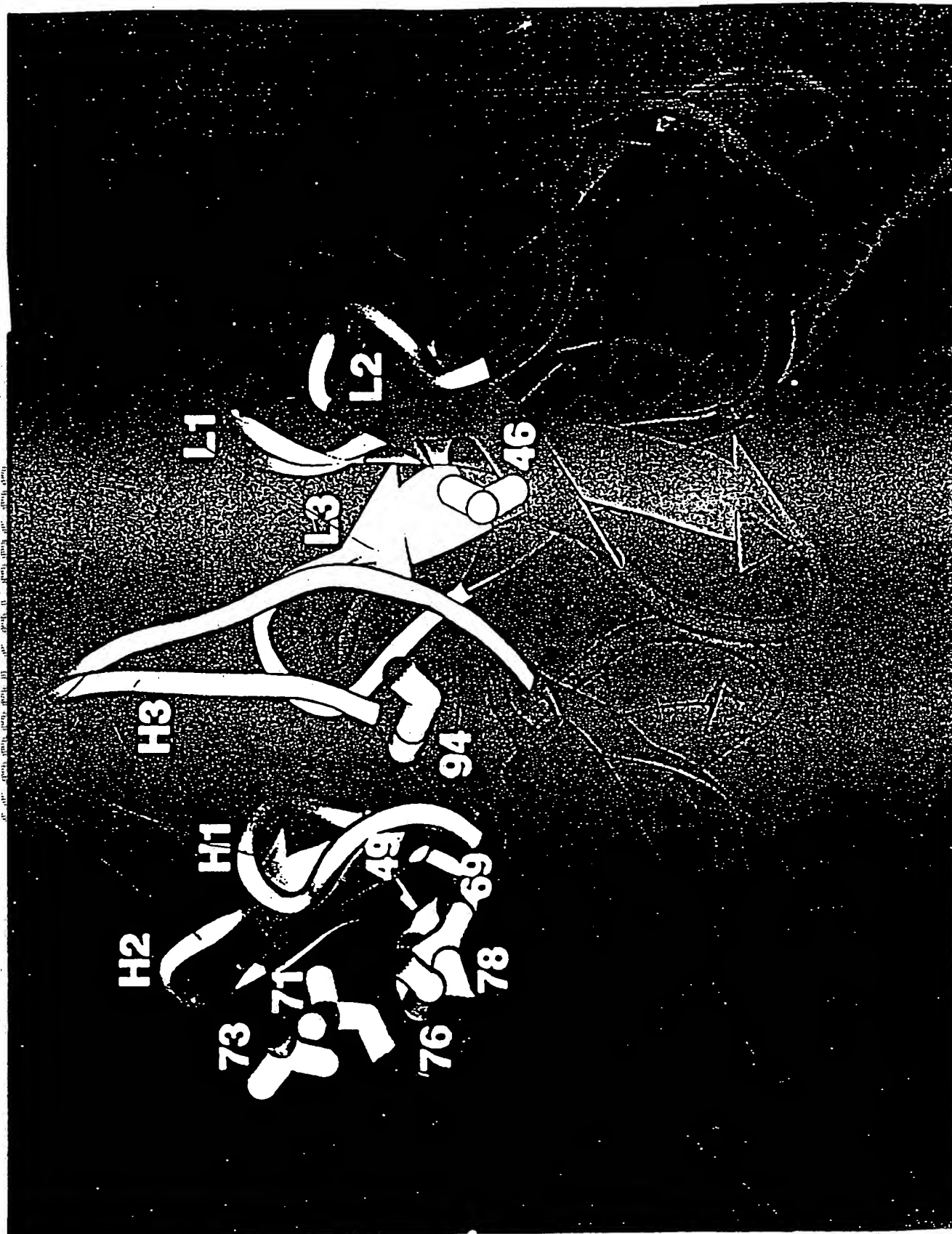


Fig. 2

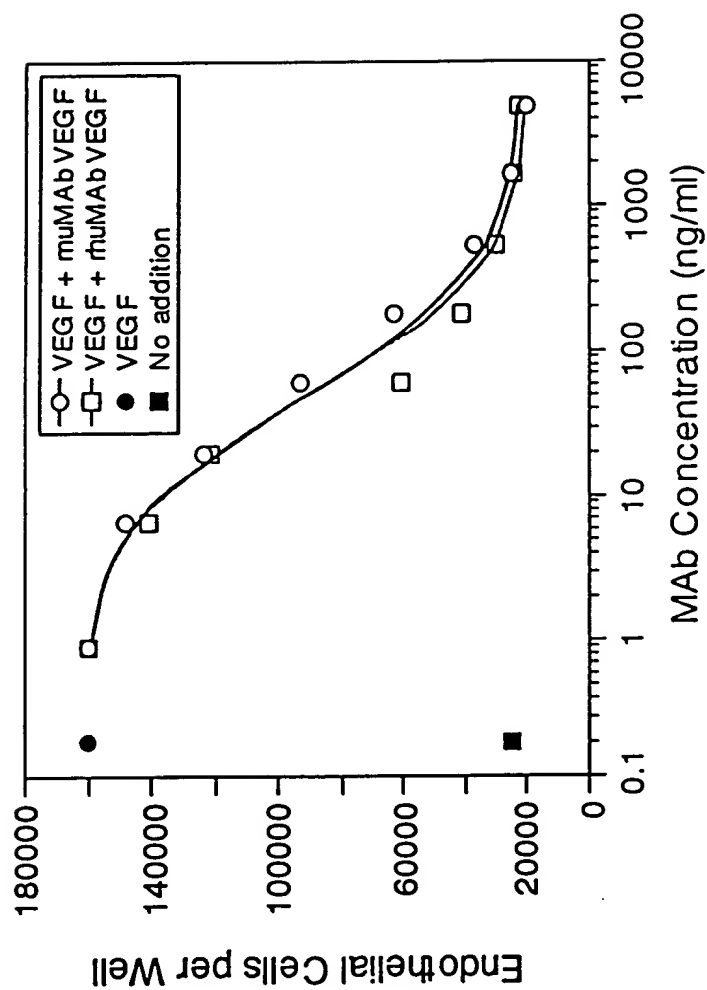


Fig. 3

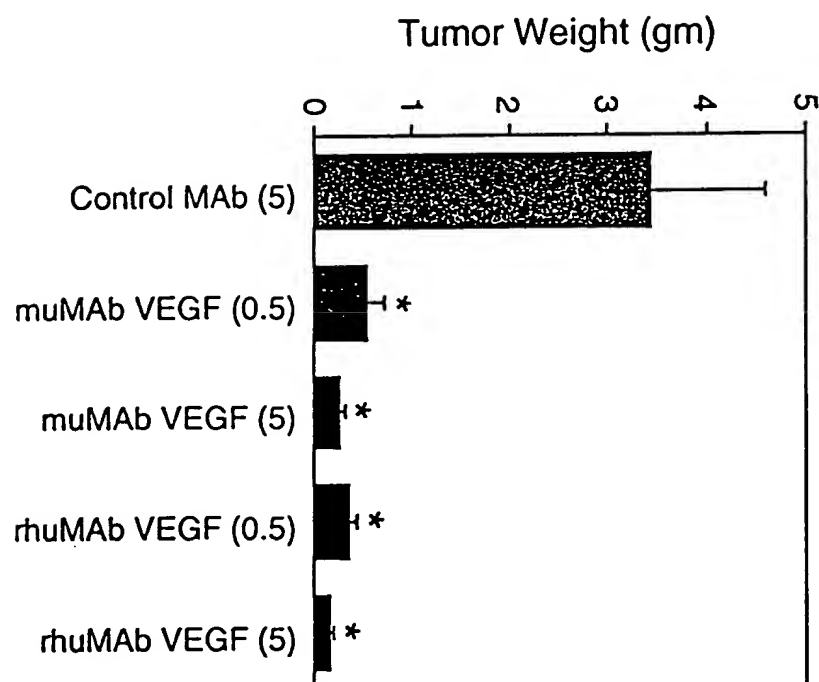


Fig. 4

V_L domain

	10	20	30	40
A4.6.1	DIQMTQTSSSLASLGDRVIISCSASQDISNYLNWYQQKP			
	** * * *			
hu2.0	DIQMTQSPSSLSASVGDRVTITCSASQDISNYLNWYQQKP			
hu2.10	DIQMTQSPSSLSASVGDRVTITCSASQDISNYLNWYQQKP			

Fig. 5A

	50	60	70	80
A4.6.1	DGTVKVLIIYFTSSLHSGVPSRFSGSGSGTDYSLTISNI·EP			
	*****		** * *	
hu2.0	GKAPKLLIYFTSSLHSGVPSRFSGSGSGTDFTLTISLQP			
hu2.10	GKAPKLLIYFTSSLHSGVPSRFSGSGSGTDYTLTISLQP			

	90	100	
A4.6.1	EDIATYYCQYSTVPWTFGGGKLEIK (SEQ ID NO:10)		
	* * *		
hu2.0	EDFATYYCQYSTVPWTFGQGTKVEIK (SEQ ID NO:13)		
hu2.10	EDFATYYCQYSTVPWTFGQGTKVEIK (SEQ ID NO:15)		

- V_H domain

	10	20	30	40
A4.6.1	EIQLVQSGPELKQPGETVRISCKASGYTFTNYGMNWVKQA			
	* * * * * * * *			*
hu2.0	EVQLVESGGGLVQPGGSLRLSCAASGYTFTNYGMNWVRQA			
hu2.10	EVQLVESGGGLVQPGGSLRLSCAASGYTFTNYGMNWIRQA			

Fig. 5B

	50 a	60	70	80
A4.6.1	PGKGLKWMGWINTYTGEPTYAADFKRRFTFSLETSASTAYL			
	* *		* * * * *	
hu2.0	PGKGLEWVGWINTYTGEPTYAADFKRRFTISRDNKNTLYL			
hu2.10	PGKGLEWVGWINTYTGEPTYAADFKRRFTISLDTSASTVYL			

	abc	90	100abcdef	110
A4.6.1	QISNLKNDdTATYFCAKYPHYYGSSHWYFDVWGAGTTVTVSS (SEQ ID NO:9)			
	*** *** * * *		* *	
hu2.0	QMNSLRAEDTAVYYCARYPHYYGSSHWYFDVWGQGLTVTVSS (SEQ ID NO:14)			
hu2.10	QMNSLRAEDTAVYYCAKYPHYYGSSHWYFDVWGQGLTVTVSS (SEQ ID NO:16)			

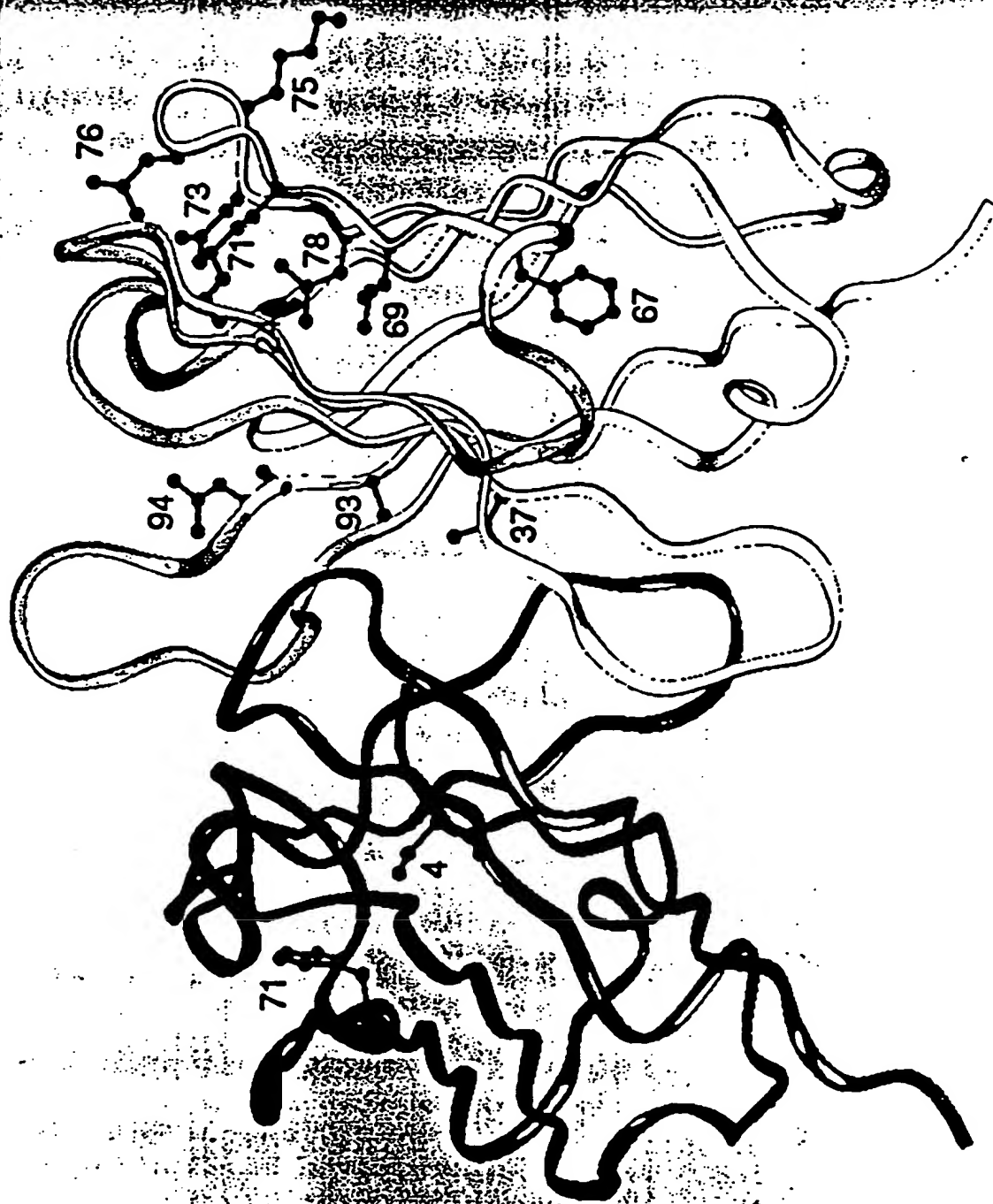
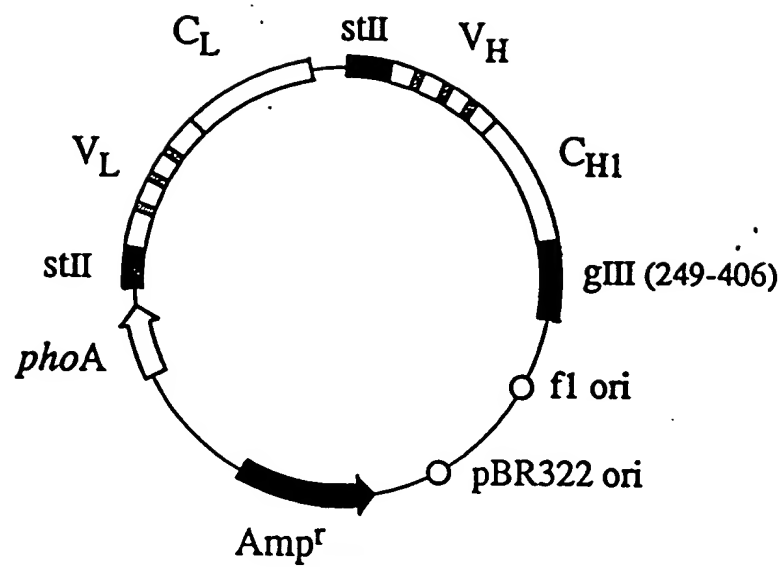


Fig. 6



Transform *E. coli*

+ M13KO7 helper phage

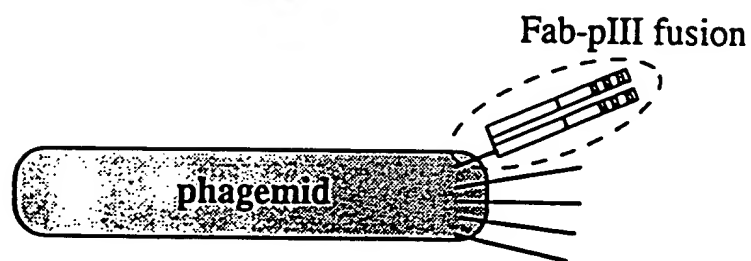


Fig. 7

1 GAATTCAACT TCTCCATACT TTGGATAAGG AAATACAGAC ATGAAAAATC TCATTGCTGA GTTGTATTATTT AAGCTTTTGA GATTATCGTC ACTGCAATGC
CTTAAGTTGA AGAGGTATGA AACCTATTCC TTTATGTCTG TACTTTTTAG AGTAACGACT CAACAATAAA TTCGAAACCT CTAATAGCAG TGACGTTACG

101 TTCGCAATAT GCGGCAAAAT GACCAACAGC GGTGATTGA TCAGGTAGAG GGGGCGCTGT ACAGGATAA GCCCGATGCC AGCATTCCTG ACAGCATATC
AAGCGTTATA CCGCGTTTGA CTGGTTGTCG CCAACTAACT AGTCCATCTC CCGCGGACA TGCTCCATTI CCGGCTACGG TCGTAAGGAC TGCTGCTATG

201 GGAGCTGCTG CGCATATTACG TAAAGAAGTT ATTGAAGCAT CCTCGTCAGT AAAAAGTTAA TCTTTTCAAC AGCTGTCTATA AAGTTGTACAC GGCGGAGACT
CCTCGACGAC GCGCTAATGC ATTTCTTCAA TAACTTCGTA GGAGCAGTCA TTTTCAATT AGAAAAAGTTG TCGACAGTAT TTCAACAGTG CCGGCTCTGA

301 TATAGTCGCT TTGTTTTTAT TTTTAAATGT ATTTGTAATCT AGAATTCGAG CTCGCTACCC GGGGATCCTC TAGAGGTTGA GGTGATTTTA TGA AAAAAGAA
ATATCAGCGA AACAAAAATA AAAAATTACA TAAACATTGA TCTTAAGCTC GAGCCATGGG CCCCTAGGAG ATCTCCAAT CCACTAAAAT ACTTTTCTT
M etLysLysAsn
-23 Begin stII signal sequence

401 TATCGCATTT CTTCTTGCAT CTATGTTCTGT TTTTCTTATT GCTACAAAACG CGTACGCTGA TATCCAGTTG ACCAGTCCC CGAGCTCCCT GTCCGCTCT
ATAGCGTAAA GAAGAACGTA GATACAAGCA AAAAAGATAA CGATGTTTGC GCATGCGACT ATAGGTCAAC TGGGTCAGGG GCTCGAGGGA CAGGCGGAGA

-19 IleAlaPhe LeuLeuAlaSer erMetPheVa lPheSerIle AlaThrAsnA lATyrAlaAs pileGlnLeu ThrGlnSerP roSerSerLe userAlaSer
Begin light chain

501 GTGGGCGATA GGGTCACCAT CACCTGCAGC GCAAGTCAGG ATATTAGCAA CTATTAAAC TGGTATCAAC AGAAACCAGG AAAAGCTCCG AAACTACTGA
CACCCGCTAT CCCAGTGGTA GTGGACGTCG CGTTCAGTCC TATAATCGTT GATAAATTG ACCATAGTTG TCTTTGGTCC TTTTCGAGGC TTTGATGACT

15 ValGlyAspA rgValThrIl eThrCysSer AlaSerGlnA spIleSerAs nTyrLeuAsn TrpTyrGlnG lNlysProGl yLysAlaPro LysLeuLeuIle

601 TTTACTTCAC CTCCTCTCTC CACTCTGGAG TCCCTTCTCG CTTCTCTGGA TCCGGTTCTG GGACGGATTA CACTCTGACC ATCAGCAGTC TGCAGCCAGA
AAATGAAGTG GAGGAGAGAG GTGAGACCTC AGGGAAGAGC GAAGAGACCT AGGCAAGAC CTGCTCTAAT GTGAGACTGG TAGTCGTGAG ACGTCGGTCT

49 TyrPheTh rSerSerLeu HisSerGlyV alProSerAr gPheSerGly SerGlySerG lyThrAspTy rThrLeuThr ileSerSerL euGlnProGlu

701 AGACTTCGCA ACTTATTACT GTCAACAGTA TAGCACCGTG CCGTGGACGT TTGGACAGGG TACCAAGGTG GAGATCAAAAC GAACGTGTGGC TGCACCATCT
TCTGAAGCGT TGAATAATGA CAGTTGTGAT ATCGTGGCAC GGCACCTGCA AACCTGTCCC ATGTTTCCAC CTCTAGTTTG CTTGACACCG ACGTGGTAGA

82 AspPheAla ThrTyrTyrC ysGlnGlnTy rSerThrVal ProTrpThrP heGlyGlnG lYThrLysVal GluIleLysA rgThrValAl aAlaProSer

801 GTCTTCATCT TCCCGCCATC TGATGAGCAG TTGAAATCTG GAACGTCTTC TGTGTGTGTC CTGCTGAATA ACTTCTATCC CAGAGAGGCC AAAGTACAGT
CAGAAAGTAGA AGGCGGTAG ACTACTCGTC AACCTTTAGAC CTTGACGAAG ACAACACACG GACGACTTAT TGAAGATAGG GTCTCTCCGG TTTCATGTCA

115 ValPheIleP heProProSe rAspGluGln LeuLysSerG lyThrAlaSe rValValCys LeuLeuAsnA snPheTyrPr oArgGluAla LysValGlnTrp

901 GGAAGGTGGA TAACGCCCTC CAATCGGGA ACTCCAGGA GAGTGTACA GAGCAGGACA GCAAGGACAG CACCTACAGC CTCAGCAGCA CCTGACGCT
CCTTCCACCT ATTGCGGAG GTTAGCCCAT TGAGGTCCT CTACACAGTG CTGCTCTGT CGTTCCTGTG GTGATGTGCG GAGTCGTCTG GGGACTGCGA

149 LysValas pasnAlaLeu GlnSerGlyA snSerGlnG lUserValThr GluGlnAspS erLysAspSe rThrTyrSer LeuSerSerT hrLeuThrLeu

1001 GAGCAAGCA GACTACGAGA AACACAAAGT CTACGCCCTG GAAGTCACCC ATCAGGGCCT GAGCTCGCCC GTCACAAAAGA GCTTCAACAG GGGAGAGTGT
CTCGTTTCTG CTGATGCTCT TTGTGTTTCA GATGCGGACG CTTCAAGTGG TAGTCCCGGA CTCGAGCGGG CAGTGTCTTCT CGAAGTTGTC CCTCTCACA

182 SerLysAla AspTyrGluL ysHisLysVa lTyrAlaCys GluValThrH isGlnGlyLe uSerSerPro ValThrLysS erPheAsnAr gGlyGluCys

Fig. 8A

1101 TAAGCTGATC CTCTACGCCG GACGCATCGT GGCCTAGTA CGCAACTAGT CGTAAAGG GATCTAGAG GTTGAGGTGA TTTTATGAAA AAGAATATCG
 ATTCGACTAG GAGATCGGC CTGCGTAGCA CCGGGATCAT CGGTGATCA GCATTTTCC CATAGATCTC CAACTCCACT -23 MetLys LysAsnIleAla
 215 OC* Begin stII signal sequence

1201 CATTCTTCT TGCATCTATG TTCTGTTTTT CTATTGCTAC AAACGCGTAC GCTGAGGTTC AGCTGGTGA GTCTGGCGGT GGCCTGGTGC AGCCAGGGGG
 GTAAAGAAGA ACGTAGATAC AAGCAAAAAA GATAACGATG TTTGCGCATG CGACTCCAAG TCGACCACT CAGACCGCCA CCGGACCACG TCGGTCCCCC
 -17 PheLeuLe uAlaSerMet PheValPheS erIleAlaTh rAsnAlaTyr AlaGluValG InLeuValG1 userGlyGly GlyLeuValG InProGlyGly
 1301 CTCACCTCCGT TTGCTCTGTG CAGCTTCTGG CTATACCTTC ACCAACTATG GTATGAACCTG GATCCGTACG CCCCCGGTA AGGGCCTGGA ATGGGTTGGA
 GAGTGAGGCA AACAGGACAC GTCGAAGACC GATATGGAAG TGGTTGATAC CATACTTGAC CTAGGAGTC CCGGGCCCAT TCCCGGACCT TACCCAACCT
 17 SerLeuArg LeuSerCysA laAlaSerG1 yTyrThrPhe ThrAsnTyrG lYMetAsnTr pileArgGln AlaProGlyL ySGlyLeuG1 utrPvalGly
 1401 TGGATTAAACA CCTATACCGG TGAACCGACC TATGCTGCGG ATTTCAAACG TCGTTTTACT ATATCTGCAG ACACCTCCAG CAACACAGTT TACCTGCAGA
 ACCTAATTGT GGATATGGCC ACTTGCTGG ATACGACGCC TAAAGTTTGC AGCAAAATGA TATAGACGTC TGTGAGGTC GTTGTGTCAA ATGGACGTCT
 50 TrpIleAsnT hrTyrThrG1 yGluProThr TyrAlaAlaA spPheLysAr gArgPheThr ileSerAlaA spThrSerSe rAsnThrVal TyrLeuGlnMet
 1501 TGAACAGCCT GCGCGCTGAG GACACTGCCG TCTATTACTG TGCAAAGTAC CCGCACTATT ATGGGAGCAG CCACTGGTAT TTCGACGTCT GGGGTCAAGG
 ACTTGTCGGA CCGCGGACTC CTGTGACGGC AGATAATGAC ACGTTTCATG GCGGTGATAA TACCCTCGTC GGTGACCATA AAGCTGCAGA CCCCAGTTCC
 84 AsnSerLe uArgAlaGlu AspThrAlav alTyrTyrCy salaLysTyr ProHisTyrT yRGlySerSe rHisTrpTyr PheAspValt rpGlyGlnGly
 1601 AACCTGGTC ACCGTCTCCT CGGCCTCCAC CAAGGGCCCA TCGGTCTTCC CCCTGCCACC CTCCTCCAAG AGCACTCTG GGGGCACAGC GGCCCTGGGC
 TTGGGACCAAG TGGCAGAGGA GCGGAGGTG GTTCCCGGT AGCCAGAAG GGGACCGTG GAGGAGTTC TCGTGGAGC CCCCCTGTGC CCGGACCCG
 117 ThrLeuVal ThrValSers erAlaSerTh rLysGlyPro SerValPheP roLeuAlaPr oSerSerLys SerThrSerG lyGlyThrAl aAlaLeuGly
 1701 TGCCTGGTCA AGGACTACTT CCCCGAACCG GTGACGGTGT CGTGGAACCTC AGCGCCCTG ACCAGCGCG TGCACACCTT CCCGGCTGTC CTACAGTCTT
 ACGGACCAGT TCCTGATGAA GGGCTTGGC CACTGCCACA GCACCTGAG TCCCGGGAC TGGTCCCGC ACGTGTGAA GGGCCGACAG GATGTCCAGA
 150 CysLeuVal lYsAspTyrPh eProGluPro valThrValS erTrpAsnSe rGlyAlaLeu ThrSerGlyV alHisThrPh eProAlaVal LeuGlnSerSer
 1801 CAGGACTCTA CTCCTCAGC AGCGTGGTGA CCGTGCCCTC CAGCAGCTTG GGCACCCAGA CCTACATCTG CAACGTGAAT CACAAGCCCA GCAACACCAA
 GTCCTGAGAT GAGGGAGTCG TCGCACCACT GGCACGGGAG GTCGTGAAC CCGTGGGTCT GGATGTAGAC GTTGCACTTA GTGTTGGGT CGTTGTGGTT
 184 GlyLeuTy rSerLeuSer SerValValt hrValProse rSerSerLeu GlyThrGlnT hrTyrIleCy sAsnValAsn HisLysProS erAsnThrLys
 1901 GGTGACACAAG AAAGTTGAGC CCAATCTTG TGACAAAACCT CACCTCTAGA GTGGCGGTG CTCTGGTTCC GGTGATTTTG ATTATGAAA GATGGCAAAC
 CCAGCTGTTT TTTCAACTCG GGTTTAGAAC ACTGTTTGA GTGGAGATCT CACCGCCACC GAGACCAAGG CCACATAAAC TAATACTTTT CTACCGTTTG
 217 ValAspLys LysValGluP roLysSerCy sAspLysThr HisLeuAM*S erGlyGlyG1 ySerGlySer GlyAspPheA spTyrGluLy sMetalAsn
 2001 GCTAATAAGG GGGCTATGAC CGAAAATGCC GATGAAAACG CGCTACAGTC TGACGCTAAA GGCAACCTG ATTCTGTCG TACTGATTAC GGTGCTGCTA
 CGATTATTCC CCCGATACTG GCTTTTACGG CTACTTTTGC GCGATGTCAG ACTGCGATTT CCGTTTGAAC TAAGACAGCG ATGACTAATG CCACGACGAT
 250 AlaAsnLysG lYalaMetTh rGluAsnAla AspGluAsnA laLeuGlnSe rAspAlaLys GlyLysLeuA spSerValAl aThrAspTyr GlyAlaAlaIle
 2101 TCGATGGTTT CATGGTGAC GTTTCGGGC TTGCTAATGG TAATGGTGCT ACTGGTGATT TTGCTGGCTC TAATTCCCA ATGGCTCAAG TCGGTGACGG
 AGCTACC2AA GTAACCACTG CAAAGGCCG CAAAGGCCG AACGATTACC ATTACCACGA TGACCACCTAA AACGACCGAG ATTAAGGGTT TACCGAGTTC AGCCACTGCC
 284 AspGlyPh eIleGlyAsp valSerGlyL euAlaAsnG1 yAsnGlyAla ThrGlyAspP heAlaGlySe rAsnSerGln MetalaglnV alGlyAspGly
 2201 TGATAATTCA CCTTTAATGA ATAATTTCCG TCAATATTTA CCTTCCCTCC CTCAATCGGT TGAATGTCGC CCTTTTGTCT TTAGCGCTGG TAAACCATAT
 ACTATTAGT GGAATTTACT TATTAAAGC AGTTATAAT GGAAGGAGG GAGTTAGCCA ACTTACAGCG GGAAACACAGA AATCGCGACC AATTGGTATA
 317 AspAsnSer ProLeuMeta snAsnPheAr gGlnTyrLeu ProSerLeuP roGlnSerVa lGluCysArg PropheValp heSerAlaG1 yLysProTyr

Fig. 8B

00/908, 469

2301 GAATTTTCTA TTGATTGTGA CAAAATAAAC TTATTCGGTG GTGTCTTTGC GTTCTTTTA TATGTTGCCA CCTTTATGTA TGTATTTTCT ACCTTTGCTA
 CTTAAAAGAT AACTAACACT GTTTTATTG AATAAGGCACTTACAGAACCGGCAATAGGTAATATATGAAACGGT GGAATAACAT ACATAAAGA TGCAAACGAT
 350 GluPheSerI leAspCysAs pLysileAsn LeupheArgG lyValPheAl aPheLeuLeu TyrValAlat hrPheMetTy rValPheSer ThrPheAlaAsn
 2401 ACATACTGCG TAATAAGGAG TCTTAATCAT GCCAGTTCTT TTGGCTAGCG CCGCCCTATA CCTTGTCTGC CTCCCGCGGT TGCCTCGCG TGCAATGGAGC
 384 IleLeuAr gAsnLysGlu SerOC* (SEQ ID NO: 100)
 2501 CGGGCCACCT CGACCTGAAT GGAAGCCGGC GGCACCTCGC TAACGGATTG ACCACTCCAA GAATTGGAGC CAATCAATTC TTGCGGAGAA CTGTGAATGC
 GCGCGGTGGA GCTGGACTTA CCTTCGGCGC CCGTGGAGCG ATTGCCTAAG TGGTGAGGTT CTTAACCTCG GTTAGTTAAG AACGCCTCTT GACACTTACG
 2601 GCAAAACCAAC CCTTGGCAGA ACATATCCAT CGCGTCCGCC ATCTCCAGCA GCCGCACGCG GCGCATCTCG GGCAGCGTTG GGTCTGGCC ACGGGTGCGC
 CGTTTGGTTG GGAACCGTCT TGTATAGGTA GCGCAGGCGG TAGAGGTCTG CCGCGTGGCG CCGCTGCAAC CCAGGACCGG TGCCACGCG
 2701 ATGATCGTGC TCCTGTCTGT GAGGACCCGG CTAGGCTGGC GGGGTTGCCT TACTGTTAG CAGAATGAAT CACCGATACG CGAGCGAAGC TGAAGCGACT
 TACTAGCACG AGGACAGCAA CTCCTGGGCC GATCGGACCG CCCAACGGA ATGACCAATC GTCTTACTTA GTGGCTATGC GCTCGCTTGC ACTTCGCTGA
 2801 GCTGCTGCAA AACGCTGCG ACCTGAGCAA CAACATGAAT GGTCTTCGGT TTCCGTGTTT CGTAAAGTCT GGAACCGCG AGTCAGCGC CCTGCACCAT
 CGACGACGTT TTGCAGACGC TGGACTCGTT GTTGTACTTA CCAGAGGCA AAGGCACAAA GCATTTTCTA CTTTTCGCG TTTCAGTCGG GACGTGGTA
 2901 TATGTTCCGG ATCTGCATCG CAGGATGCTG CTGGCTACCC TGTGGAACAC CTACATCTGT ATTACGAAG CGCTGGCATT GACCTTGAGT GATTTTCTC
 ATACAAGGCC TAGACGTAGC GTCCTACGAC GACCGATGGG ACACCTTGTG GATGTAGACA TAATTGCTTC GCGACCGTAA CTGGGACTCA CTAATAAAGAG
 3001 TGGTCCCGCC GCATCCATAC CGCCAGTTGT TTACCTTCAC AACGTTCCAG TAACCGGGCA TGTTTCATCAT CAGTAACCCG TATCGTGAGC ATCCTCTCTC
 ACCAGGGCGG CGTAGGTATG GCGGTCAACA AATGGGAGTG TTGCAAGGTC ATTGGCCCGT ACAAGTAGTA GTCATTTGGC ATAGCACTCG TAGGAGAGAG
 3101 GTTTCATCGG TATCATTACC CCCATGAACA GAAATTCGCC CTTACACGGA GGCATCAAGT GACCAACACG GAAAAACCG CCTTAAACAT GGCCCGCTTT
 CAAAGTAGCC ATAGTAATGG GGGTACTTGT CTTTAAAGGG GAATGTGCTT CCGTAGTTCA CTGGTTTGTG CTTTTCGCG GGAATTTGTA CCGGGCGAAA
 3201 ATCAGAAGCC AGACATTAA CTTCTCTGGAG AAACCTCAACG AGCTGGACGC GGATGAACAG GCAGACATCT GTGAATCGCT TCACGACCCAC GCTGATGAGC
 TAGCTTTCGG TCTGTAATTG CGAAGACCTC TTTGAGTTGC TCGACCTGCG CCTACTTGTG CGTCTGTAGA CACTTAGCGA AGTGCTGGTG CGACTACTCG
 3301 TTTACCGCAG GATCCGGAAA TTGTAAACGT TAATATTTG TTAAATTCG CGTTAAATTT TGTTAATC AGCTCATTTT TTAACCAATA GGCCGAAATC
 AAATGGCGTC CTAGGCCTTT AACATTGTGA ATTATAAAAC AATTITAAGC GCAATTTAAA AACCAATTTAG TCGAGTAAAA AATTGGTTAT CCGGCTTTAG
 3401 GGCAAAATCC CTTATAAATC AAAAGAATAG ACCGAGATAG GGTGAGTGT TGTCCAGTT TGGAAACAAGA GTCCACTATT AAAGAACGCTG GACTCCAACG
 CCGTTTATAG GAATATTTAG TTTTCTTATC TGGCTCTATC CCAACTCACA ACAAGGTCAA ACCTTGTCTT TGGAAACAAGA GTCCACTATT AAAGAACGCTG GACTCCAACG
 3501 TCAAGGGCGG AAAAACCGTC TATCAGGGCT ATGGCCCACT ACGTGAACCA TCACCCCTAAT TCACCTTTT GGGGTCGAGG TGCCGTAAAG CACTAAATCG
 AGTTTCCCGC TTTTGGCAG ATAGTCCCGA TACCGGGTGA TGCACTTGGT AGTGGATTA GTTCAAAAAA CCCCAGCTCC ACGGCATTTT GTGATTTAGC
 3601 GAACCCCTAA GGGAGCCCC GATTTAGAGC TTGACGGGGA AAGCCGGCGA ACGTGGCGAG AAAGGAAGGG AAGAAAGCGA AAGGAGCGGG CGCTAGGGCG
 CTTGGGATTT CCCTCGGGGG CTAATCTCG AACTGCCCTT TTCGGCCGCT TGCACCGCTC TTTCTTCC TTTCTTCCGT TTCTTCGCG CCGATCCCGC
 3701 CTGGCAAGTG TAGCGGTCAC GCTGCGCGTA ACCACCACAC CCGCCCGGCT TAATGCCCGG CTACAGGGCG CGTCCGGATC CTGCCCTCGG CGTTTCGGTG
 GACCGTTTAC ATCGCCAGTG CGACGCGCAT TGGTGGTGTG GCGCGCGCG ATTACCGGCG GATGTCCCG GACGGAGCGC GCAAAGCCAC
 3801 ATGACGGTGA AAACCTCTGA CACATGCAGC TCCCGGAGAC GGTACAGCT TGTCTGTAAG CCGATGCCGG GAGCAGACAA GCGCGTCAGG GCGCGTCAGC
 TACTGCCACT TTTGGAGACT GTGTACGTG AGGCCCTCTG CCAGTGTGCA ACAGACATTC GCCTACGGCC CTCGTCTGTT CCGGCGAGTCC CCGCGAGTCC

Fig. 8C

3901 GGGTGTGGC GGGTGTGGG GGCAGGCCAT GACCCAGTCA CGTAGCGATA GCGGAGTGA TACTGGCTTA ACTATGCGGC ATCAGAGCAG ATTGTACTGA
CCCACAACCG CCCACAGCCC CGGTGCGGTA CTGGGTGAGT GATGGCTAT GCGCTGACAT ATGACCGAAT TGATACGCCG TAGTCTCGTC TAACATGACT
4001 GAGTGCACCA TATGCGGTGT GAAATACCGC ACAGATGCGT AAGGAGAAAA TACCGCATCA GCGGCTCTTC CGCTTCCTCG CTCACTGACT CGCTGCGCTC
CTCACGTGGT ATACGCCACA CTTTATGGCG TGCTACGCA TTCCTCTTTT ATGGCGTAGT CCGCGAGAAG GCGAAGGAGC GAGTGACTGA GCGACCGCAG
4101 GGTGCTTCGG CTGCGGCGAG CCGTATCAGC TCACTCAAAG GCGGTAATAC GGTATCCAC AGAATCAGGG GATAACGCAG GAAAGAACAT GTGAGCAAAA
CCAGCAAGCC GACGCGGCTC GCCATAGTCG AGTGAGTTTC CGCCATTATG CCAATAGGTG TCTTAGTCCC CTATTGCGTC CTTTCTTGTA CACTCGTTTT
4201 GGCCAGCAAA AGCCAGGAA CCGTAAAAAG GCGCGTTTC TGGCGTTTTT CCATAGGCTC CGCCCCCTG ACAGAGCATCA CAAAAATCGA CGCTCAGTC
CCGTGCTTT TCGGTCTT GGCATTTTC CGGCGCAACG ACCGCAAAA GGTATCCGAG GCGGGGGAC TGCTCGTAGT GTTTTAGCT GCGAGTTTCAG
4301 AGAGGTGGC AAACCGACA GGAATATAA GATACAGGC GTTCCCTT GGAAGCTCCC TCGTGGCTC TCCTGTTCCG ACCCTGCCG TTACCGGATA
TCTCCACCGC TTGCGCTGT CCGATATTT CTATGGTCCG CAAAGGGGA CTTTCGAGG AGCACGGAG AGGACAAGG TGGACGGC AATGGCCTAT
4401 CCTGTCCGC TTCTCCCTT CCGGAAGCGT GCGCTTTCT CATAGCTCAC GCTGTAGGTA TCTCAGTTTC GTGTAGTTCG TCGCTCCAA GCTGGGCTGT
GGACAGGCGG AAAGAGGGA GCCCTTCGA CCGGAAAGA GTATCGAGTG CGACATCCAT AGAGTCAAGC CACATCCAGC AAGCGAGGT CGACCCGACA
4501 GTGCACGAAC CCCCCTTCA GCGGACCGC TCGCCTTAT CCGGTAACTA TCGTCTTGAG TCCAAACCCG TAAGACACGA CTTATCGCCA CTGGCAGCAG
CACGTGCTTG GGGGCAAGT CCGGCTGGC ACGCGGAATA GGCCATTGAT AGCAGAACTC AGGTGGGCC ATTCTGTGCT GAATAGCGGT GACCGTCGTC
4601 CCACTGGTAA CAGGATTAGC AGAGCGAGGT ATGTAGGCGG TGCTACAGAG TTCTTGAAGT GGTGGCCTAA CTACGGCTAC ACTAGAAGGA CAGTATTGG
GGTGACCAT GTCTAATCG TCTCGTCCA TACATCCGCC ACGATGTCTC AAGAACTTCA CCACCGGATT GATGCCGATG TGATCTTCT GTCATNAACC
4701 TATCTGCGT CTGCTGAAGC CAGTTACCTT CGGAAAAAGA GTTGGTAGCT CTTGATCCG CAAACAACC ACCGCTGGTA GCGGTGGTTT TTTTGTGTTG
ATAGACCGGA GACGACTCG GTCAATGGAA GCCTTTTCT CAACCATCGA GAACCTAGGC GTTGTGTTGG TGGCGACCAT CGCCACCAA AAAACAAACG
4801 AAGCAGCAGA TTACGCGCAG AAAAAAGGA TCTCAAGAAG ATCCTTTGAT CTTTCTACG GGTCTGACG CTCAGTGGAA CGAAAACTCA CGTTAAGGGA
TTGCTGCTCT AATGCGGCTC TTTTCTTCT AGAGTTCTTC TAGGAACTA GAAAGATGC CCCAGACTGC GAGTCACTT GCTTTGAGT GCAATTCCCT
4901 TTTTGGTCAAT GAGATTATCA AAAAGGATCT TCACCTAGAT CCTTTAAAT TAAAAATGA GTTTAAATC AATCTAAAGT ATATATGAGT AAACCTGGTC
AAAACCACTA CTCTAATAGT TTTTCTCTAGA AGTGGATCTA GAAAAATTA ATTTTACTT CAAAATTTAG TTAGATTTC TATATACTCA TTTTGAACCCAG
5001 TGACAGTTAC CAATGCTTAA TCAGTGAGGC ACCTATCTCA GCGATCTGTC TATTTCTGTC ATCCATAGTT GCTGACTCC CCGTCTGTA GATAACTACG
ACTGTCAATG GTTACGAAT AGTCACTCCG TGGATAGAGT CGCTAGACAG ATAAAGCAAG TAGGTATCAA CCGACTGAG GGCAGCACAT CTATTGATGC
5101 ATACGGGAGG GCTTACCATC TGGCCCCAGT GCTGCAATGA TACCGCGAGA CCCACGCTCA CCGSCTCCAG ATTTATCAGC AATAAACCCAG CCAGCCGGAA
TATGCCCTCC CGAATGGTAG ACCGGGTCA CGACGTTACT ATGGCGCTCT GGTGCGAGT GCGCGAGGTC TAAATAGTCG TTATTTGGTC GGTGCGCCTT
5201 GGGCCGAGCG CAGAAGTGGT CCTGCAACTT TATCGCCTC CATCCAGTCT ATTAATTGTT GCCGGGAAGC TAGAGTAAGT AGTTCGCCAG TTAATAGTTT
CCCGGCTCGC GTCTTACCA GACGTTGAA ATAGGCGGAG GTAGGTGAGA TAATTAACA CGGCCCTTCG ATCTCATTC TCAAGCGGTC AATTATCAAA
5301 GCGCAACGTT GTTGCCATTG CTGCAGGCAT CGTGGTGTC GCGTCTGCT TGGTATGGC TTCAATTCAGC TCCGGTTCCC AACGATCAAG GCGAGTTACA
CGCGTTGCAA CAACGGTAAC GACGTCCGTA GCACACAGT GCGAGCAGCA AACCATACCG AAGTAAGTCG AGGCCAAGG TTGCTAGTTC CGCTCAATGT
5401 TGATCCCCCA TGTGTGCAA AAAAGCGGTT AGCTCCTTCG GTCTCCGAT CGTTGTGAGA AGTAAGTTGG CCGCAGTGT ATCACTCATG GTTATG3CAG
ACTAGGGGCT ACAACACGTT TTTTCGCCAA TCGAGGAAGC CAGGAGGCTA GCAACAGTCT TCATTCAACC GCGGTCACAA TAGTGAGTA CAATACCGTC
5501 CACTGCATAA TTCTCTTACT GTCATGCCAT CCGTAAGATG CTTTCTGTG ACTGGTGAGT ACTCAACCAA GTCATTCTGA GAATAGTGA TCGGGCGACC
GTGACGTATT AAGAGAATGA CAGTACGGTA GGCATTCTAC GAAAAGACAC TGACCACTCA TGAGTTGGTT CAGTAAGACT CTTATCACAT ACGCCGCTGG

Fig. 8D

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0 2

5601 GAGTTGCTCT TGCCCGGGCT CAACACGGGA TAATACCGGC CGCATAGCA GAACCTTAAA AGTCTCATC ATTGGAAAAC GTTCTTCGGG GCGAAAACATC
CTCAACGAGA ACGGGCCGCA GTTGCCCT ATATGCGC GGTGATCGT CTTGAAATTT TCACGAGTAG TAACCTTTTG CAAGAAGCCC CGCTTTTGAG

5701 TCAAGGATCT TACCGCTGT GAGATCCAGT TCGATGTAAC CCACTCGTGC ACCCAACTGA TCTTCAGCAT CTTTACTTT CACCAGCGTT TCTGGGTGAG
AGTTCCTAGA ATGGCGACAA CTCTAGGTCA AGCTACATTG GGTGAGCAGC TGGGTTGACT AGAAGTCGTA GAAAATGAAA GTGGTCGCAA AGACCCCACTC

5801 CAAAAACAGG AAGGCAAAAT GCCGCAAAA AGGGAATAAG GCGACACCG AAATGTTGAA TACTCATACT CTTCCCTTTT CAATATTATT GAAGCATTTA
GTTTTTGTC TTCCGTTTTC CGGCGTTTTT TCCCTTATC CCGCTGTGCC TTTACAACCT ATGAGTATGA GAAGGAAAA GTTATAATAA CTTCGTAAAT

5901 TCAGGGTTAT TGTCTCATGA GCGGATACAT ATTTGAATGT ATTTAGAAAA ATAAACAAAT AGGGGTTCG CGCACATTC CCCGAAAAGT GCCACCTGAC
AGTCCCAATA ACAGAGTACT CGCTATGTA TAACTTTACA TAAATCTTTT TATTGTTTA TCCCAAGGC GCGTGTAAG GGGCTTTTCA CGGTGGACTG

6001 GTCTAAGAAA CCATTATTAT CATGACATTA ACCTATAAAA ATAGGCGTAT CACGAGGCCC TTTCGCTCTC AA (SEQ ID NO: 99)
CAGATTCTTT GGTAATAATA GTACTGTAAT TGGATATTTT TATCCGCATA GTGCTCCGGG AAAGCAGAAG TT

Fig. 8E

■ = differences from F(ab)-12

10 20 30
F(ab)-12 DIQMTQSPSSLSASVGDRVITITCSASQDISNYLNWYQQ
Y0243-1 DIQ■TQSPSSLSASVGDRVITITC■AN■Q■SNYLNWYQQ
Y0238-3 DIQ■TQSPSSLSASVGDRVITITC■AN■Q■SNYLNWYQQ
Y0313-1 DIQ■TQSPSSLSASVGDRVITITC■AN■Q■SNYLNWYQQ
Y0317 DIQ■TQSPSSLSASVGDRVITITCSASODISNYLNWYQQ

CDR-L1

40 50 60 70
F(ab)-12 KPGKAPKVLIIYFTSSLHSGVPSRFSGSGSGTDFTLTIS
Y0243-1 KPGKAPKVLIIYFTSSLHSGVPSRFSGSGSGTDFTLTIS
Y0238-3 KPGKAPKVLIIYFTSSLHSGVPSRFSGSGSGTDFTLTIS
Y0313-1 KPGKAPKVLIIYFTSSLHSGVPSRFSGSGSGTDFTLTIS
Y0317 KPGKAPKVLIIYFTSSLHSGVPSRFSGSGSGTDFTLTIS

Fig. 10A

CDR-L2

80 90 100
F(ab)-12 SLQPEDFATYYCQOYSTVPWTFGQGTKVEIKRTV (SEQ ID NO: 8)
Y0243-1 SLQPEDFATYYCQOYSTVPWTFGQGTKVEIKRTV (SEQ ID NO: 109)
Y0238-3 SLQPEDFATYYCQOYSTVPWTFGQGTKVEIKRTV (SEQ ID NO: 111)
Y0313-1 SLQPEDFATYYCQOYSTVPWTFGQGTKVEIKRTV (SEQ ID NO: 113)
Y0317 SLQPEDFATYYCQOYSTVPWTFGQGTKVEIKRTV (SEQ ID NO: 115)

CDR-L3

10 20 30
F(ab)-12 EVQLVESGGGLVQPGGSLRLSCAASGYTFTNYGMNWVR
Y0243-1 EVQLVESGGGLVQPGGSLRLSCAASGY■FT■YGMNWVR
Y0238-3 EVQLVESGGGLVQPGGSLRLSCAASGYTFTNYG■NWVR
Y0313-1 EVQLVESGGGLVQPGGSLRLSCAASGY■FT■YGMNWVR
Y0317 EVQLVESGGGLVQPGGSLRLSCAASGY■FT■YGMNWVR

CDR-H1

40 50 60 70
F(ab)-12 QAPGKGLEWVGWINTYTGEPTYAADFKRRFTFSLDTSKSTA
Y0243-1 QAPGKGLEWVGWINTYTGEPTYAADFKRRFTFSLDTSKSTA
Y0238-3 QAPGKGLEWVGWINTYTGEPTYAADFKRRFTFSLDTSKSTA
Y0313-1 QAPGKGLEWVGWINTYTGEPTYAADFKRRFTFSLDTSKSTA
Y0317 QAPGKGLEWVGWINTYTGEPTYAADFKRRFTFSLDTSKSTA

Fig. 10B

CDR-H2

80 90 100 110 CDR-7
F(ab)-12 YLQMNSLRAEDTAVYYCAKYPHYGYGSSHWFYFDVWGQGT (SEQ ID NO: 7)
Y0243-1 YLQMNSLRAEDTAVYYCAKYPHYGYGSSHWFYFDVWGQGT (SEQ ID NO: 110)
Y0238-3 YLQMNSLRAEDTAVYYCAKYP■YGYG■SHWFYFDVWGQGT (SEQ ID NO: 112)
Y0313-1 YLQMNSLRAEDTAVYYCAKYP■YGYG■SHWFYFDVWGQGT (SEQ ID NO: 114)
Y0317 YLQMNSLRAEDTAVYYCAKYP■YGYG■SHWFYFDVWGQGT (SEQ ID NO: 116)

CDR-H3

0303459.030697

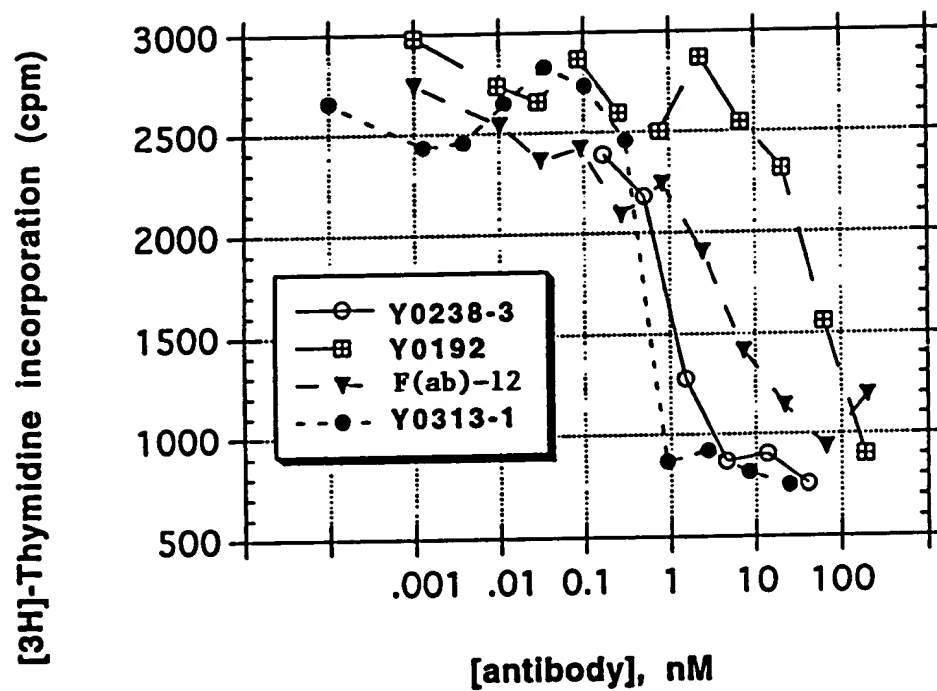


Fig. 11

Fig. 12

